

### Amendments to the Claims

**1. (Original)** A cobalamin derivative

- (a) having no binding affinity or low binding affinity to transcobalamin II and
- (b) retaining activity as a vitamin B12 substitute.

**2. (Original)** The cobalamin derivative according to claim 1

- (a) having less than 20% of binding affinity to transcobalamin II when compared to the binding affinity of non-modified cobalamin in a binding test, and
- (b) retaining more than 2% of the activity as a vitamin B12 substitute in a growth assay.

**3. (Original)** The cobalamin derivative according to claim 1

- (a) having less than 10% of binding affinity to transcobalamin II when compared to the binding affinity of non-modified cobalamin in a binding test, and
- (b) retaining more than 10% of the activity as a vitamin B12 substitute in a growth assay.

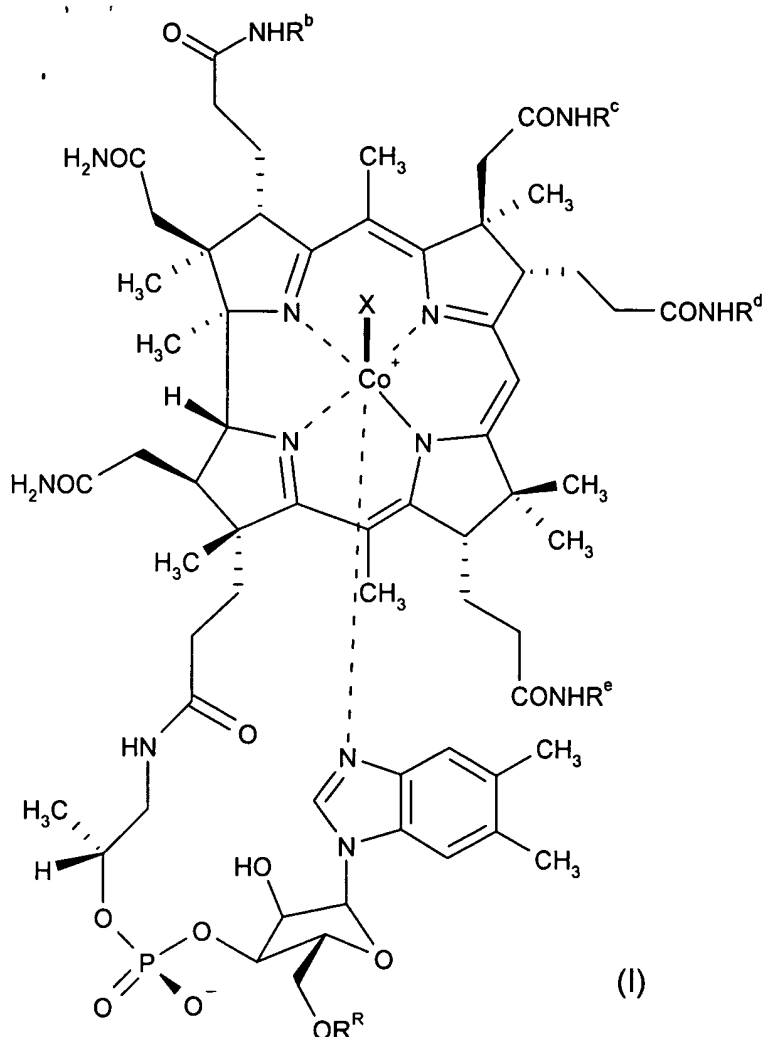
**4. (Original)** The cobalamin derivative according to claim 1

- (a) having less than 5% of binding affinity to transcobalamin II when compared to the binding affinity of non-modified cobalamin in a binding test, and
- (b) retaining more than 10% of the activity as a vitamin B12 substitute in a growth assay.

**5. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 1 to 4~~ claim 1 carrying a therapeutic and/or diagnostic agent.

**6. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 1 to 5~~ claim 1 carrying a radioactive metal.

**7. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 1 to 6~~ claim 1 of formula (I)



wherein

$R^b$ ,  $R^c$ ,  $R^d$  and  $R^e$ , independently of each other, are a spacer-chelator group, an antibiotic or antiproliferative therapeutic agent, a sterically demanding organic group with 4 to 20 carbon atoms, or hydrogen;

$R^R$  is a spacer-chelator group or an antibiotic or antiproliferative therapeutic agent, each connected through a linker Z, or hydrogen;

with the proviso that at least three of the residues  $R^b$ ,  $R^c$ ,  $R^d$ ,  $R^e$  and  $R^R$  are hydrogen and at least one of the residues  $R^b$ ,  $R^c$ ,  $R^d$  and  $R^e$  is different from hydrogen;

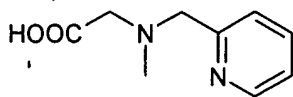
X is a monodentate ligand; and

the central cobalt (Co) atom is optionally in the form of a radioactive isotope.

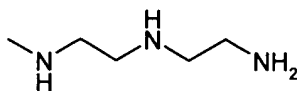
**8. (Original)** The cobalamin derivative according to claim 7 wherein R<sup>e</sup> is hydrogen.

**9. (Currently amended)** The cobalamin derivative according to claim 7 ~~or 8~~ wherein the spacer-chelator group comprises  
a spacer, which is an aliphatic chain of 2 to 10 carbon atoms, wherein one or two carbon atoms may be replaced by nitrogen and/or oxygen atoms and the aliphatic chain may be substituted by hydroxy, oxo or amino, and  
a chelator, which is a compound having two, three or more donor atoms selected from nitrogen, oxygen and sulfur in a distance such as to bind to a metal atom, and optionally a metal atom.

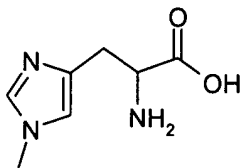
**10. (Original)** The cobalamin derivative according to claim 9 wherein the chelator is selected from the chelators of formula (II) to (IX),



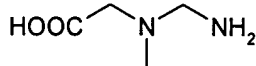
(II)



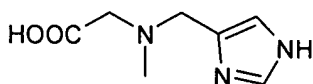
(III)



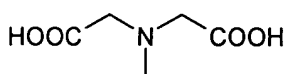
(IV)



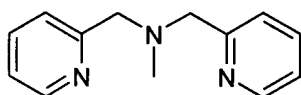
(V)



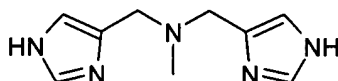
(VI)



(VII)



(VIII)



(IX)

wherein carboxyl groups may be present as esters.

**11. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 6 to 10~~ claim 6 wherein the radioactive metal is  $^{94m}\text{Tc}$ ,  $^{99m}\text{Tc}$ ,  $^{188}\text{Re}$ ,  $^{186}\text{Re}$ ,  $^{111}\text{In}$ ,  $^{90}\text{Y}$ ,  $^{64}\text{Cu}$ ,  $^{67}\text{Cu}$  or  $^{177}\text{Lu}$ .

**12. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 7 to 11~~ claim 7 wherein X is cyano, methyl, hydroxy, aquo or a 5'-deoxyadenosyl group.

**13. (Original)** The cobalamin derivative according to claim 12 wherein X is cyano.

**14. (Currently amended)** The cobalamin derivative according to ~~anyone of claims 7 to 12~~ claim 7 wherein the central cobalt atom is the radioisotope  $^{57}\text{Co}$  or  $^{60}\text{Co}$ .

**15. (Original)** The cobalamin derivative according to claim 10 wherein  $R^b$  is a spacer-chelator group optionally carrying a metal atom, the spacer is an aliphatic chain of 2 to 4 carbon atoms, and the chelator is of formula (II), wherein the group COOH is optionally in the form of an ester;  $R^c$ ,  $R^d$ ,  $R^e$  and  $R^R$  are hydrogen; and X is cyano.

**16. (Original)** The cobalamin derivative according to claim 15 wherein  $R^b$  is a spacer-chelator group optionally carrying a metal atom, the spacer is an aliphatic chain of 4 carbon atoms, and the chelator is of formula (II), wherein the group COOH is in the form of the ethyl ester;  $R^c$ ,  $R^d$ ,  $R^e$  and  $R^R$  are hydrogen; and X is cyano.

**17. (Original)** The cobalamin derivative according to claim 10 wherein  $R^d$  is a spacer-chelator group optionally carrying a metal atom, the spacer is an aliphatic chain of 3 carbon atoms, and the chelator is of formula (II), wherein the group COOH is optionally in the form of an ester;  $R^b$ ,  $R^c$ ,  $R^e$  and  $R^R$  are hydrogen; and X is cyano.

**18. (Original)** The cobalamin derivative according to claim 10 wherein  $R^b$  is a spacer-chelator group optionally carrying a metal atom, the spacer is an aliphatic chain of 2 carbon atoms, and the chelator is of formula (III);  $R^c$ ,  $R^d$ ,  $R^e$  and  $R^R$  are hydrogen; and X is cyano.

**19. (Currently amended)** A pharmaceutical composition comprising a cobalamin derivative according to ~~anyone of claims 1 to 18~~ claim 1.

**20. (Currently amended)** A method of diagnosis of a neoplastic disease or an infection by microorganisms in a mammal comprising

- (a) exposing the mammal suspected of being inflicted by a neoplastic disease or an infection to a period of a vitamin B12 – free diet, and
- (b) subsequently applying a cobalamin derivative according to ~~anyone of claims 1 to 18~~ claim 1 carrying a diagnostic agent.

**21. (Currently amended)** A method of treatment of a mammal suffering from a neoplastic disease or an infection by microorganisms comprising

- (a) exposing the mammal in need of treatment to a period of a vitamin B12 – free diet, and
- (b) subsequently applying a cobalamin derivative according to ~~anyone of claims 1 to 18~~ claim 1 carrying a therapeutic agent.

**22-25. (Cancelled)**

**26. (New)** The method of claim 20, wherein the cobalamin is effective in cancer imaging.

**27. (New)** The method of claim 21, wherein the cobalamin is effective in cancer imaging.